AMENDMENTS TO THE CLAIMS:

This listing of claims will replace all prior versions, and listings of claims in the application:

LISTING OF CLAIMS:

1.(original) A transgenic non-human animal expressing at least one transgene comprising a DNA sequence encoding a heterologous Amyloid Precursor Protein(APP) comprising at least the Arctic mutation (E693G) and a further AD (Alzheimer's disease) pathogenic mutation or a further transgene affecting AD pathogenesis, which results in increased amounts of intracellular soluble Aβ aggregates, including Aβ peptides.

2.(original) The transgenic animal according to claim 1, wherein the transgene/transgenes are integrated in the genomic DNA.

3.(currently amended) The transgenic animal according to claim 1 [[or 2]], wherein said transgene/transgenes are operably linked to a promoter effective for expression of said gene in the brain tissue of said animal.

4.(currently amended) The transgenic animal according to ~~any of claims 1-3~~ claim 1, wherein the endogenous APP is expressive or non-expressive.

5.(currently amended) The transgenic animal according to ~~any of claims 1-4~~ claim 1, wherein said further transgene is a human presenilin-1 and/or presenilin-2 transgene harboring an AD pathogenic mutation.

6.(currently amended) The transgenic animal according to ~~any of claims 1-4~~ claim 1, wherein said further transgene comprises a DNA sequence encoding apolipoprotein E, apolipoprotein J(clusterin), a_1-antichymotrypsin (ACT) or fragments thereof.

7.(currently amended) The transgenic animal according to ~~any of claims 1-4~~ claim 1, wherein said further AD pathogenic mutation is one of the APP mutations KM670/671DF, KM670/671DY, KM670/671EF or KM670/671EY.

8.(currently amended) The transgenic animal according to ~~any of claims 1-4~~ claim 1, wherein said further AD pathogenic mutation is one of the APP

mutations KM670/671NL, KM670/671NY, KM670/671NF, KM670/671KL, KM670/671DL or KM670/671EL, wherein KM670/671NL (the Swedish mutation) is preferred.

9.(currently amended) The transgenic animal according to ~~any of claims 1-4~~ claim 1, wherein the transgenic animal expresses only one transgene which comprises only the Arctic mutation (E693G) and the Swedish mutation (KM670/671NL).

10.(currently amended) The transgenic animal according to ~~any of claims 1-9~~ claim 1, additionally comprising a homologously integrated targeting construct for at least one of the neprilysin or insulin-degrading enzyme (IDE) genes, which disrupts these genes through gene ablation (knock-out) and enhances Aβ-40 and/or Aβ-42 Arctic peptide production.

11.(currently amended) The transgenic animal according to ~~any of claims 1-10~~ claim 1, wherein the transgenic animal is a rodent.

12.(currently amended) The transgenic animal according to ~~any of claims 1-11~~ claim 1, wherein the transgenic animal is a murine animal.

13.(currently amended) The transgenic animal according to ~~claim 1-12~~ claim 1, wherein the transgenic animal is a mouse.

14.(currently amended) A method of producing the transgenic animal according to ~~any of claims 1-13~~ claim 1, comprising integrating in the genomic DNA at least one transgene comprising a DNA sequence encoding a heterologous Amyloid Precursor Protein (APP) comprising at least the Arctic mutation (E693G) and a further AD (Alzheimer's disease) pathogenic mutation or a further transgene affecting AD pathogenesis.

15.(original) The method according to claim 14, wherein said transgene/transgenes are operably linked to a promoter effective for expression of said gene in the brain tissue of said animal.

16.(currently amended) The method according to ~~any of claims 14-15~~ claim 14, wherein the endogenous APP is optionally made non-expressive.

17.(currently amended) The method according to ~~any of claims 14-16~~ claim 14, wherein said further transgene is a human presenilin-1 and/or presenilin-2 transgene harboring an AD pathogenic mutation.

18.(currently amended) The method according to ~~any of claims 14-16~~ claim 14, wherein said further transgene comprises a DNA sequence encoding apolipoprotein E, apolipoprotein J (clusterin), al-antichymotrypsin (ACT) or fragments thereof.

19.(currently amended) The method according to ~~any of claims 14~~ claim 14, wherein said further AD pathogenic mutation is one of the APP mutations KM670/671DF, KM670/671DY, KM670/671EF or KM670/671EY.

20.(currently amended) The method according to ~~any of claims~~ claim 14, wherein said further AD pathogenic mutation is one of the APP mutations KM670/671NL, KM670/671NY, KM670/671NF, KM670/671KL, KM670/671DL or KM670/671EL, wherein KM670/671NL (the Swedish mutation) is preferred.

21.(currently amended) The method according to ~~any of claims 14-20~~ claim 14, additionally comprising homologously integrating a targeting construct for at least one of the neprilysin or insulin-degrading enzyme (IDE) genes.

22.(currently amended) A method of screening, wherein the transgenic animal according to ~~any of claims 1-13~~ claim 1 is used for screening for agents useful for treating, preventing or inhibiting Alzheimer's disease.

23.(currently amended) A method of screening, wherein the transgenic animal according to ~~any of claims 1-13~~ claim 1 is used for screening for diagnostic agents for Alzheimer's disease.